FIGURE 1

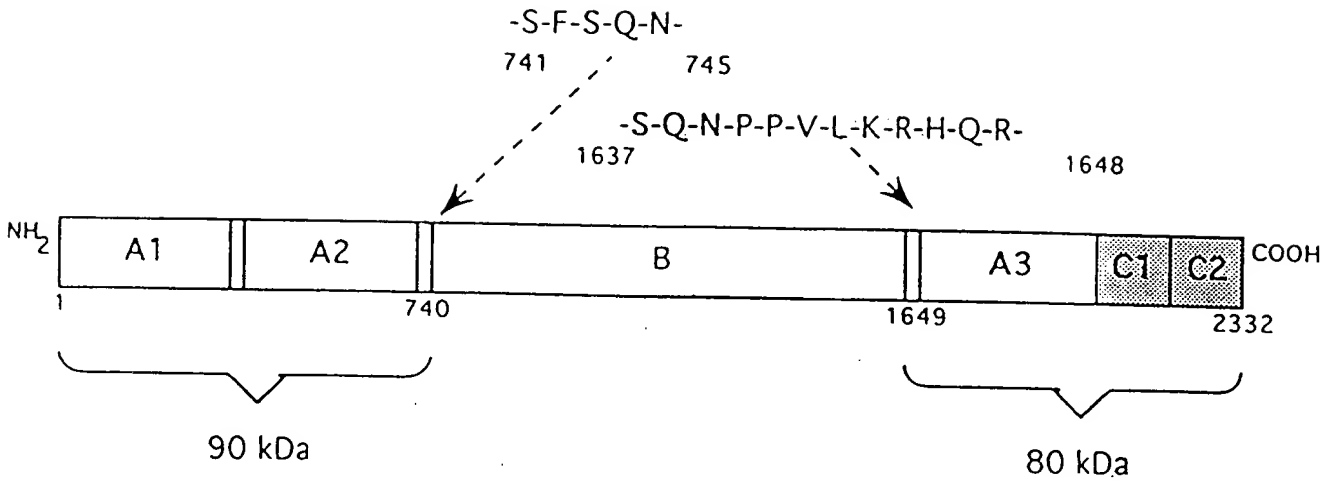
-S-F-S-Q-N-
741
745
-S-Q-N-P-P-V-L-K-R-H-Q-R-
1637
1648
NH2
A1
A2
B
A3
C1
C2
COOH
1
740
1649
2332
90 kDa
80 kDa

FIGURE 2

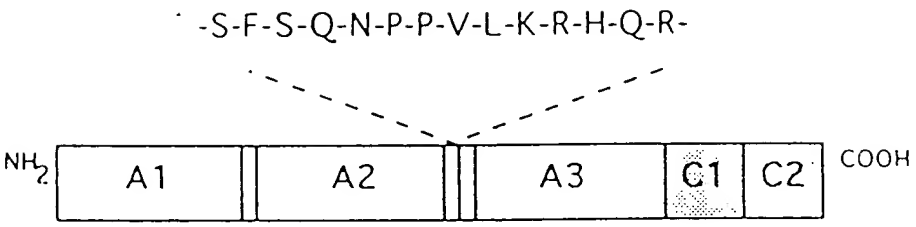

FIGURE 3

AAV-F8-1

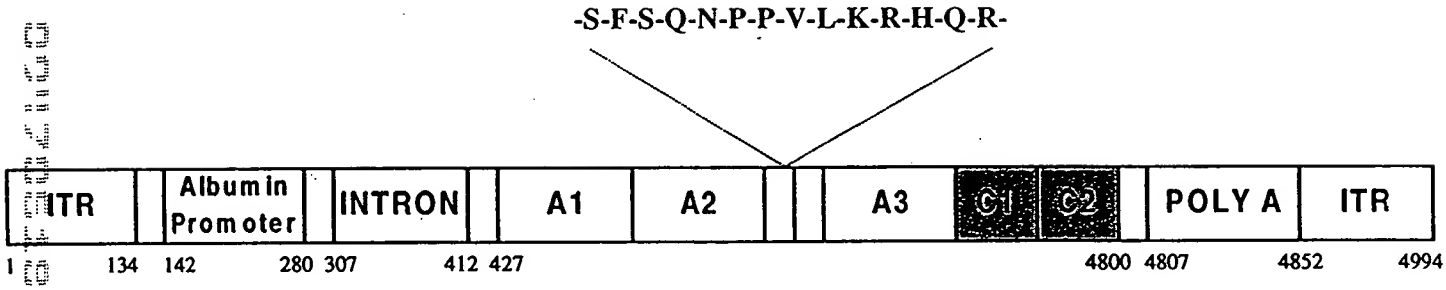

FIGURE 4

PVM4.1c-F8AB

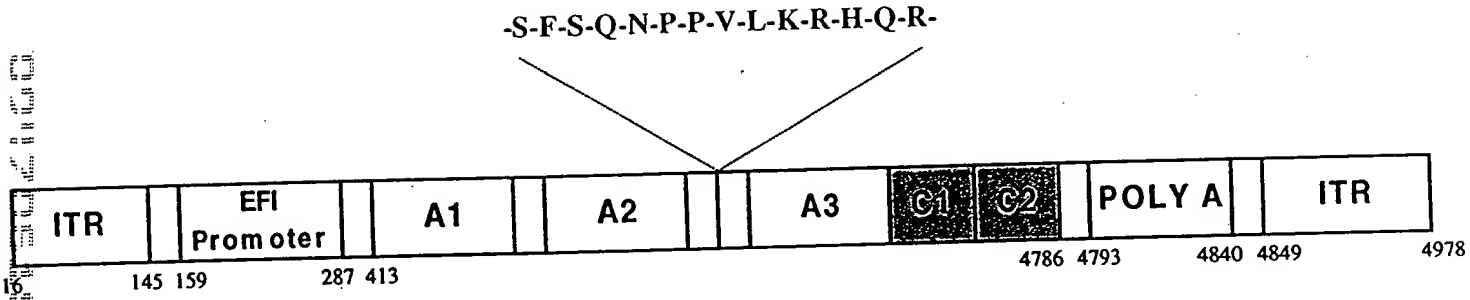

FIGURE 5

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FIGURE 6

CGCCCCTGCAGGCAGCTGCGCGCTCGCTCGCTCACTGAGGCCGCCCGGGCAA
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CTGCTCAGCATGGGCAGCAATGAAAACATCCATTCTATTCATTTCAGTGGACA
TGTGTTCACTGTACGAAAAAAAGAGGAGTATAAAATGGCACTGTACAATCTC
TATCCAGGTGTTTTTGAGACAGTGGAAATGTTACCATCCAAAGCTGGAATTTG
GCGGGTGGAATGCCTTATTGGCGAGCATCTACATGCTGGGATGAGCACACTT
TTTCTGGTGTACAGCAATAAGTGTCAGACTCCCCTGGGAATGGCTTCTGGACA
CATTAGAGATTTTCAGATTACAGCTTCAGGACAATATGGACAGTGGGCCCCA
AAGCTGGCCAGACTTCATTATTCCGGATCAATCAATGCCTGGAGCACCAAGG
AGCCCTTTTCTTGGATCAAGGTGGATCTGTTGGCACCAATGATTATTCACGGC
ATCAAGACCCAGGGTGCCCGTCAGAAGTTCTCCAGCCTCTACATCTCTCAGTT
TATCATCATGTATAGTCTTGATGGGAAGAAGTGGCAGACTTATCGAGGAAAT
TCCACTGGAACCTTAATGGTCTTCTTTGGCAATGTGGATTCATCTGGGATAAA
ACACAATATTTTTAACCCTCCAATTATTGCTCGATACATCCGTTTGCACCCAA
CTCATTATAGCATTCGCAGCACTCTTCGCATGGAGTTGATGGGCTGTGATTTA
AATAGTTGCAGCATGCCATTGGGAATGGAGAGTAAAGCAATATCAGATGCAC
AGATTACTGCTTCATCCTACTTTACCAATATGTTTGCCACCTGGTCTCCTTCAA
AAGCTCGACTTCACCTCCAAGGGAGGAGTAATGCCTGGAGACCTCAGGTGAA
TAATCCAAAAGAGTGGCTGCAAGTGGACTTCCAGAAGACAATGAAAGTCACA
GGAGTAACTACTCAGGGAGTAAAATCTCTGCTTACCAGCATGTATGTGAAGG
AGTTCCTCATCTCCAGCAGTCAAGATGGCCATCAGTGGACTCTCTTTTTTCAG
AATGGCAAAGTAAAGGTTTTTCAGGGAAATCAAGACTCCTTCACACCTGTGG

TGAACTCTCTAGACCCACCGTTACTGACTCGCTACCTTCGAATTCACCCCCAG
AGTTGGGTGCACCAGATTGCCCTGAGGATGGAGGTTCTGGGCTGCGAGGCAC
AGGACCTCTACTGACTCGAGCCTAATAAAGGAAATTTATTTTCATTGCAATAG
TGTGTTGGTTTTTTGTGTGCGGCCGCAGGAACCCCTAGTGATGGAGTTGGCCA
CTCCCTCTCTGCGCGCTCGCTCGCTCACTGAGGCCGGGCGACCAAAGGTCGCC
CGACGCCCGGGCTTTGCCCGGGCGGCCTCAGTGAGCGAGCGAGCGCGCAGCT
GCCTGCAGGACAT

FIGURE 7

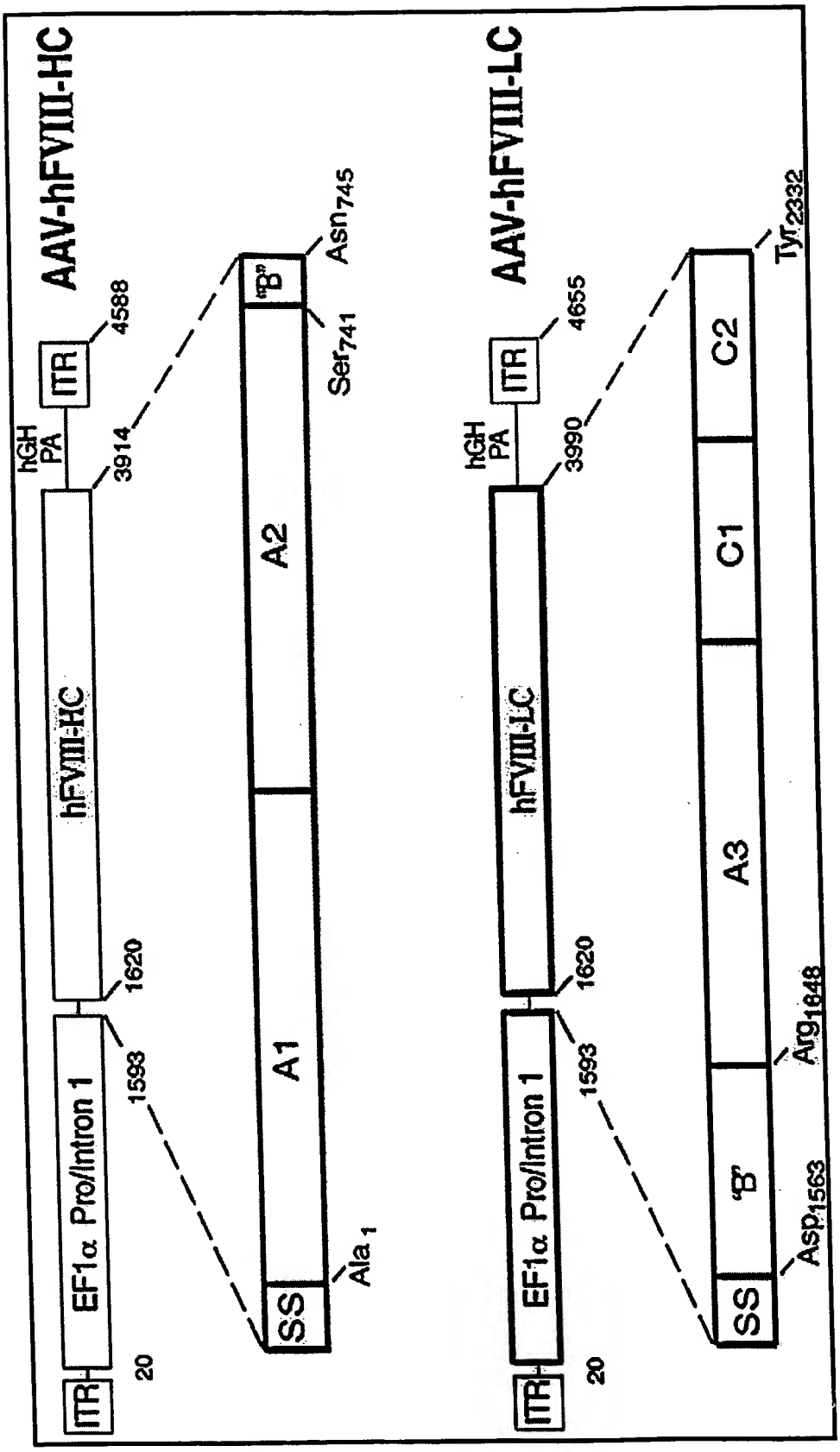

AAV-hFVIII-HC
ITR
20
EF1α Pro/Intron 1
1593
hFVIII-HC
1620
hGH PA
3914
ITR
4588
SS
A1
A2
"B"
Ala 1
Ser741
Asn745
AAV-hFVIII-LC
ITR
20
EF1α Pro/Intron 1
1593
hFVIII-LC
1620
hGH PA
3990
ITR
4655
SS
"B"
A3
C1
C2
Asp1563
Arg1648
Tyr2332

FIGURE 8

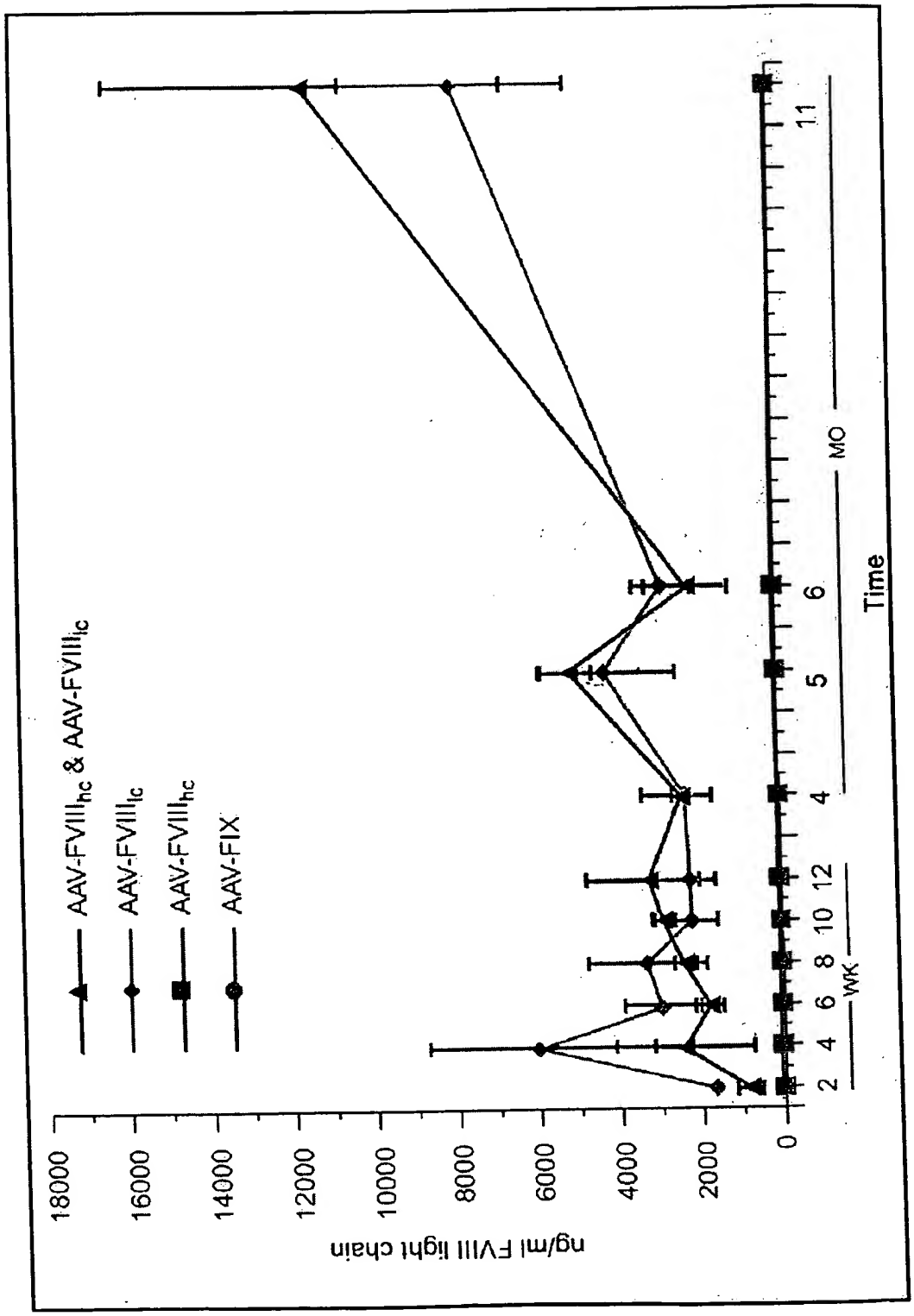

AAV-FVIII$_{hc}$ & AAV-FVIII$_{lc}$
AAV-FVIII$_{lc}$
AAV-FVIII$_{hc}$
AAV-FIX
ng/ml FVIII light chain
18000
16000
14000
12000
10000
8000
6000
4000
2000
0
2 4 6 8 10 12
WK
4 5 6
MO
11
Time

FIGURE 9

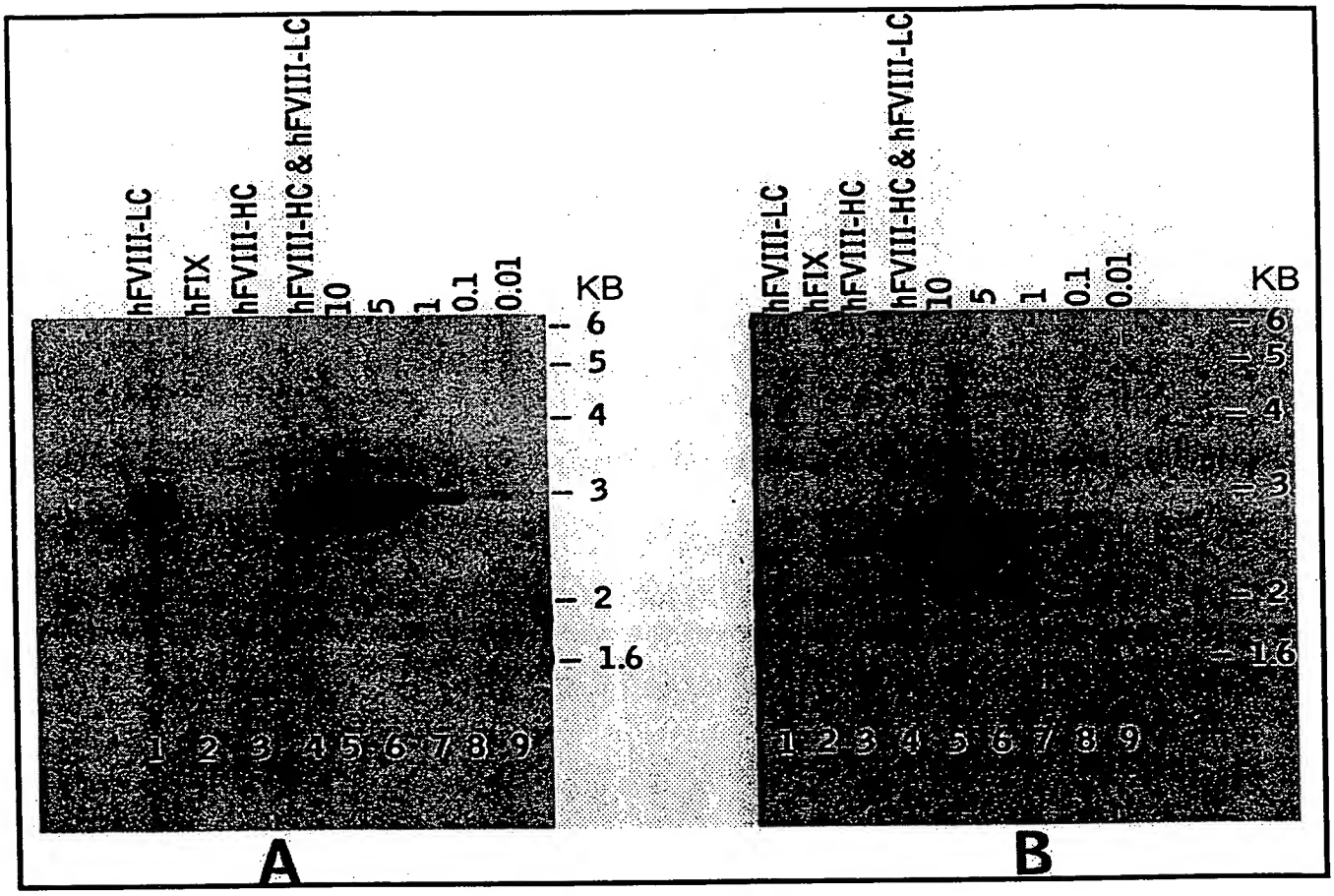

hFVIII-LC
hFIX
hFVIII-HC
hFVIII-HC & hFVIII-LC
10
5
1
0.1
0.01
KB
6
5
4
3
2
1.6
1 2 3 4 5 6 7 8 9
A
B

FIGURE 10

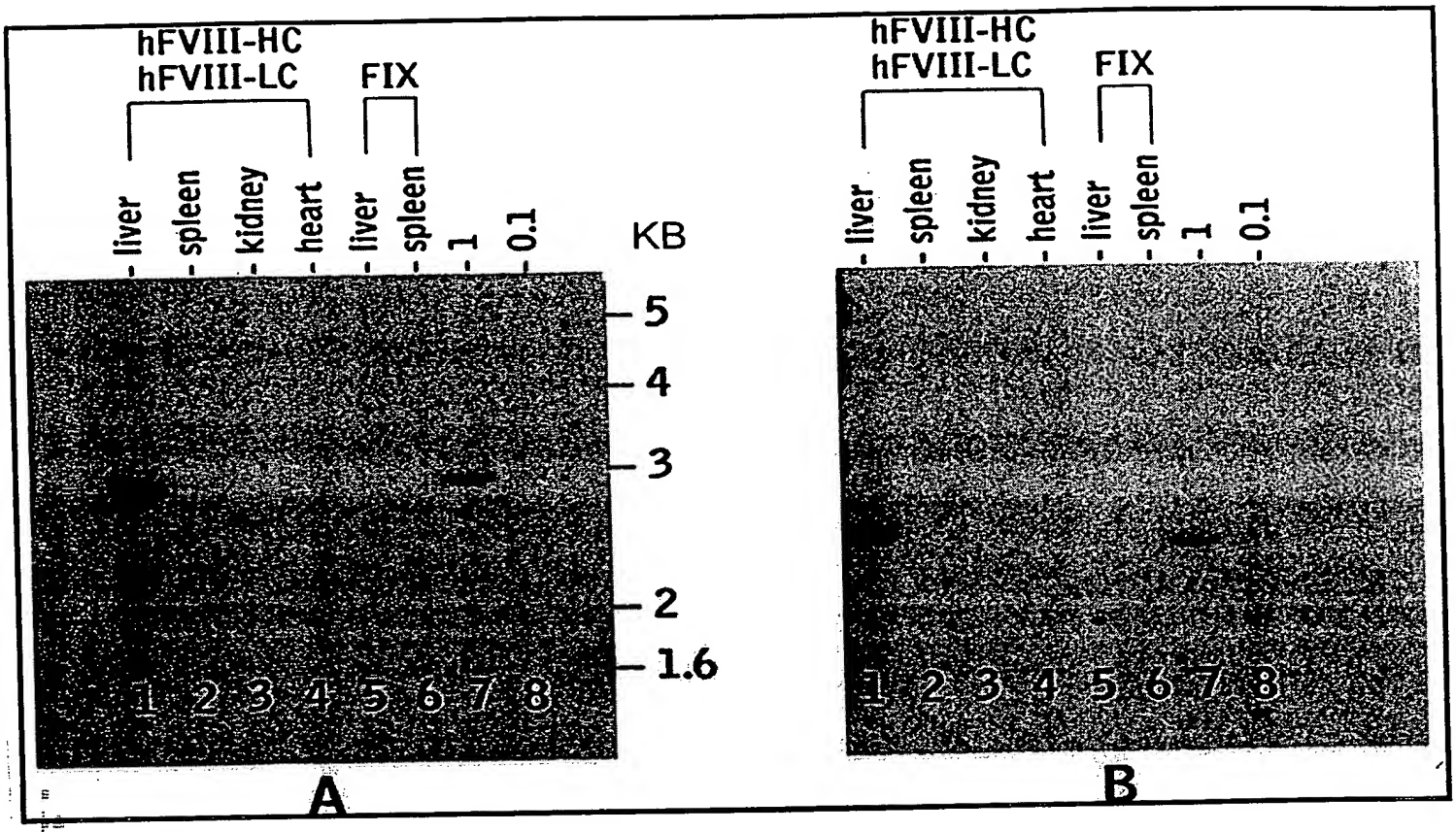
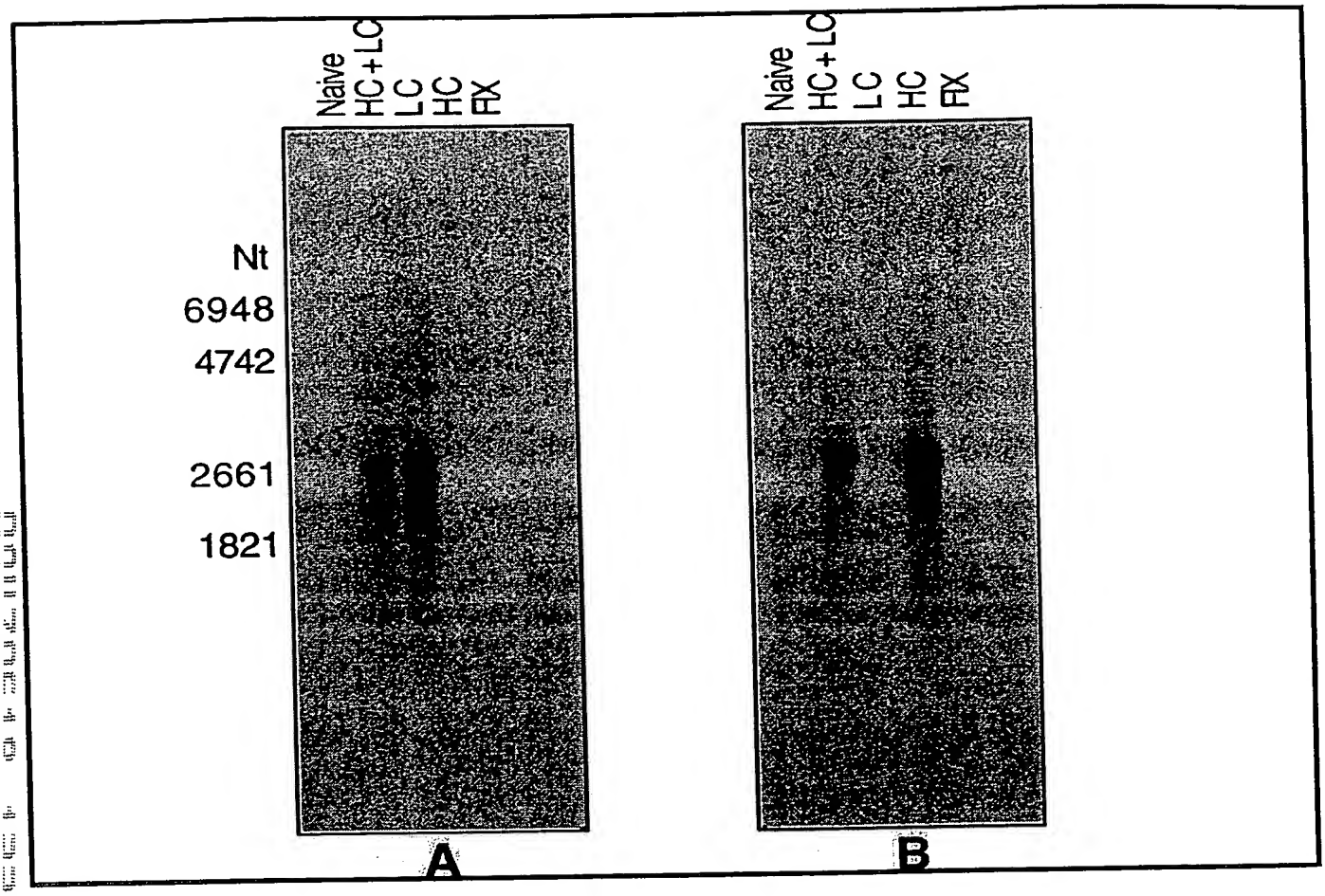

hFVIII-HC
hFVIII-LC
FIX
liver
spleen
kidney
heart
liver
spleen
1
0.1
KB
5
4
3
2
1.6
1 2 3 4 5 6 7 8
A
hFVIII-HC
hFVIII-LC
FIX
liver
spleen
kidney
heart
liver
spleen
1
0.1
1 2 3 4 5 6 7 8
B

FIGURE 11


Naive
HC+LC
LC
HC
PX
Nt
6948
4742
2661
1821
A
Naive
HC+LC
LC
HC
PX
B